

**PATENT**

**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE  
(Case No. 98,164-A24)**

**In the Application of:** )  
 )  
**Andrew A. Wolff** )  
 )  
**Serial No.: To Be Assigned** )  
 )  
**Filing Date: Concurrently Herewith** )  
 )  
**For: Sustained Release Ranolazine** )  
**Formulations** )

**INFORMATION DISCLOSURE STATEMENT**

**Commissioner of Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450**

Dear Sir:

Pursuant to 37 C.F.R. Section 1.97 - 1.99, the Applicant wishes to make the following references of record in the above-identified application. This Information Disclosure Statement is in compliance with the continuing duty of candor as set forth in 37 C.F.R. Section 1.56. Pursuant to 37 C.F.R. §1.98(d), copies of references numbered 1-43 are not provided herewith, since they were previously provided in the parent cases, U.S. Patent Application Serial No. 09/925,871, filed on August 9, 2001; and U.S. Patent Application Serial No. 09/321,522 filed on May 27, 1999 now U.S. Patent No. 6,303,607 issued October 16, 2001. These references are also listed on the enclosed PTO Form 1449.

In the judgment of the undersigned, portions of the listed references may be material to the Examiner's consideration of the presently pending claims. This statement is not a representation that the listed references have effective dates early enough to be "prior art" within the meaning of 35 U.S.C. Section 102 or Section 103.

Applicants do not believe any fee is due with this submission. If this belief be in error and the Patent Office determines that the fee prescribed in the relevant portion of 37 C.F.R. Section 1.97 is applicable, the undersigned attorney by his signature hereby authorizes any such fee to be debited from Deposit Account 13-2490.

## U. S. PATENTS

1. MacFarlane, et al., U.S. Patent No. 5,209,933, issued May 11, 1993
2. Kluge, et al., U.S. Patent No. 4,567,264, issued January 28, 1986
3. Samuels, et al., U.S. Patent No. 5,472,707, issued December 5, 1995
4. Dow, et al., U.S. Patent No. 5,506,229, issued April 9, 1996
5. Santus, et al., U.S. Patent No. 5,670,171, issued September 23, 1997

## FOREIGN PATENT DOCUMENTS

6. WO Patent No. WO 94/26266, published November 24, 1994

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8. Jain D, Dasgupta P, Hughes LO, Lahiri A, Raftery EB. Ranolazine (RS-43285): A preliminary study of a new anti-anginal agent with selective effect of ischaemic myocardium. *Eur J Clin Pharmacol* 38:111-114, 1990.
9. Lodge JPA, Lam FT, Perry SL, Giles GR. Ranolazine - a new drug with beneficial effects on renal preservation. *Transplantation* 50:755-759, 1990.
10. Cocco G, Rousseau MF, Bouvy T, Cheron P, Williams G, Detry JM, Pouleur H. Effects of a new metabolic modulator, ranolazine, on exercise tolerance in angina pectoris patients treated with  $\beta$ -blocker or diltiazem. *J Cardiovasc Pharmacol* 20:131-138, 1992.
11. Allely MC, Brown CM, Kenny BA, Kilpatrick AT, Martin A, Spedding M. Modulation of  $\beta_1$ -adrenoceptors in rat left ventricle by ischaemia and acyl carnitines: protection by ranolazine. *J Cardiovasc Pharmacol* 21:869-873, 1993.
12. Clarke B, Spedding M, Patmore L, McCormack JG. Protective effects of ranolazine in guinea-pig hearts during low-flow ischaemia and their association with increases in active pyruvate dehydrogenase. *Br J Pharmacol* 109:748-750, 1993.
13. Black SC, Gralinski MR, McCormack JG, Driscoll EM, Lucchesi BR. Effect of ranolazine on infarct size in a canine model of regional myocardial ischemia/reperfusion. *J Cardiovasc Pharmacol* 24:921-928, 1994.
14. Gralinski MR, Black SC, Kilgor KS, Chou AY, McCormack JG, Lucchesi BR. Cardioprotective effects of ranolazine (RS-43285) in the isolated perfused rabbit heart. *Cardiovascular Research* 28:1231-1237, 1994.
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19. Wyatt KM, Skene C, Veitch K, Hue L, McCormack JG. The anti-anginal agent ranolazine is a weak inhibitor of the respiratory complex I, but with greater potency in broken or uncoupled than in coupled mitochondria. *Biochemical Pharmacol* 50:1599-1606, 1995.
20. Allen TJ, Chapman RA. Effects of ranolazine on L-type calcium channel currents in guinea-pig single ventricular myocytes. *Br J Pharmacol* 118:249-254, 1996.
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23. McCormack JG, Baracos VE, Barr R, Lopaschuk GD. Effects of ranolazine on oxidative substrate preference in epitrochlearis muscle. *Am Physiol Society*, 905-910, 1996.
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27. McCormack JG, et al., "Ranolazine; a novel metabolic modulator for the treatment of angina." *Gen Pharmacol*. 1998 May; 30(5): 639-45.
28. Bagger JP, et al., "Effects of ranolazine on ischemic threshold, coronary sinus blood flow, and myocardial metabolism in coronary artery disease." *Cardiovasc Drugs Ther*. 1997 Jul;11(3): 479-84.
29. Aaker A, et al., "Effects of ranolazine on the exercise capacity of rats with chronic heart failure induced by myocardial infarction." *J Cardiovasc Pharmacol*. 1996 Sept; 28(3):353-62.

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31. Alley MC, Alps BJ. The effects of the novel anti-anginal agent ranolazine (I.D.) in a canine model of transient myocardial ischaemia. Br J Pharmacol 93(Suppl):246P, 1988
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34. Pouleur H, Hue L, Harlo BJ, Rousseau MF. Metabolic pathways modulation: a new approach to treat myocardial ischemia? Circulation 80, (Supp II):52, 1989.
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36. Ferrandon, P et al. Ranolazine but not allopurinol or dipyridamole reduces the ventricular acidosis and lactate produced by low perfusion flow in isolated rat hearts. Therapie 45:10, 1990.
37. Cocco G, Rousseau MF, Bouvy T, Cheron P, William C, Detry JM, Pouleur H. Effects of a novel metabolic modulator, ranolazine, on exercise tolerance and left ventricular filling dynamics in patients with angina petoris. Eur Heart J 13(Suppl):97, 1992.
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(note: the data in this abstract are from the correct analysis.)

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*(note: the data in the abstract are not from the final analysis.)*
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Respectfully submitted,  
**McDonnell Boehnen Hulbert & Berghoff**

Date: July 7, 2003

By: 

A. Blair Hughes  
Reg. No. 32,901

FORM PTO-1449  
(Rev. 2-32)

U.S. Department of Commerce  
Patent and Trademark Office

INFORMATION DISCLOSURE  
STATEMENT BY APPLICANT  
(Use several sheets if necessary)

Atty. Docket No.

98,164-A24

Serial No.

To Be Assigned

Applicant:

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U.S. PATENT DOCUMENTS

Examiner Initial		Document Number	Date	Name	Class	Subclasses	Filing Date if Appropriate
		5,209,933	5/11/93	MacFarlane et al.			
		4,567,264	1/28/96	Kluge et al.			
		5,472,707	12/5/95	Samuels et al.			
		5,506,229	4/9/96	Dow et al.			
		5,670,171	9/23/97	Santus et al.			

FOREIGN PATENT DOCUMENTS

		Document Number	Date	Country	Class	Subclass	Translation	
							Yes	No
		WO 94/26266	11/24/94					

OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc).

			Allely MC, Alps BJ. Prevention of myocardial enzyme release by ranolazine in a primate model of ischaemia with reperfusion. Br J Pharmacol 99:5-6, 1990.
			Jain D, Dasgupta P, Hughes LO, Lahiri A, Raftery EB. Ranolazine (RS-43285): A preliminary study of a new anti-anginal agent with selective effect of ischaemic myocardium. Eur J Clin Pharmacol 38:111-114, 1990.
EXAMINER			DATE CONSIDERED

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<b>FORM PTO-1449</b> (Rev. 2-32)  <b>INFORMATION DISCLOSURE</b> <b>STATEMENT BY APPLICANT</b> (Use several sheets if necessary)	<b>U.S. Department of Commerce</b> <b>Patent and Trademark Office</b>		<b>Atty. Docket No.</b>  98,164-A24	<b>Serial No.</b>  To Be Assigned
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			Lodge JPA, Lam FT, Perry SL, Giles GR. Ranolazine - a new drug with beneficial effects on renal preservation. Transplantation 50:755-759, 1990.
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### ABSTRACTS

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